



# **STIC Search Report**

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**STIC Database Tracking Number: 179799**

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Location: 3d65 / 3c18  
Art Unit: 1655  
Friday, March 03, 2006**

**Case Serial Number: 10/698795**

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### **Search Notes**

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L19 ANSWER 1 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2006:103807 HCAPLUS

DN 144:146025

ED Entered STN: 03 Feb 2006

TI Method for purifying virus envelope by column chromatography

IN Ioka, Shinichi

PA Genomidea, Inc., Japan

SO PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DT Patent

LA Japanese

IC ICM C07K-0001/20

ICS C07K-0014/115; C12N-0007/02

CC 9-3 (Biochemical Methods)

Section cross-reference(s): 10

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO2006011580	A1	20060202	2005WO-JP13893	20050722
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
	RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			

PRAI 2004JP-0219381 A 20040727

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2006011580	ICM	C07K-0001/20
	ICS	C07K-0014/115; C12N-0007/02
	IPCI	C07K0001-20 [ICM,7]; C07K0014-115 [ICS,7]; C12N0007-02 [ICS,7]

AB A method is provided for industrially purifying the envelope of virus (e.g., Sendai virus (Hemagglutinating Virus of Japan, HVJ)). More specifically, it is intended to provide a method for purifying an inactivated virus envelope by the combined use of ion exchange chromatog. with hydrophobic chromatog. to thereby purify the envelope at a high yield while sustaining the cell fusion activity of the virus. The virus envelope thus purified is usable as a vector for transferring a biopolymer such as a gene into cells or a living body. This method is also applicable to the purification of an attenuated envelope virus.

ST virus envelope purifn ion exchange chromatog hydrophobic

IT Alcohols, uses  
 RL: TEM (Technical or engineered material use); USES (Uses)  
 (aliphatic, lower; method for purifying virus envelope by column chromatog.)

IT Cations  
 (divalent; method for purifying virus envelope by column chromatog.)

IT Virion structure  
 (envelope, attenuated, inactivated; method for purifying virus envelope by column chromatog.)

IT Virion structure  
 (envelope; method for purifying virus envelope by column chromatog.)

IT Immunoassay  
 (hemagglutination test; method for purifying virus envelope by column chromatog.)

IT Adsorption  
 Anion exchange chromatography  
 Arenavirus  
 Buffers  
 Bunyavirus  
 Classical swine fever virus  
 Coronavirus  
 Cowpox virus  
 Crimean-Congo hemorrhagic fever virus  
 Deltavirus  
 Dengue virus  
 Ebola virus  
 Feline immunodeficiency virus  
 Filovirus  
 Flavivirus  
 Fusion, biological  
 Genetic vectors  
 Hepadnaviridae  
 Hepatitis B virus  
 Hepatitis C virus  
 Hepatitis delta virus  
 Herpesviridae  
 Human  
 Human T-lymphotropic virus 1  
 Human herpesvirus  
 Human herpesvirus 4  
 Human immunodeficiency virus  
 Hydrophobic interaction chromatography  
 Influenza virus  
 Ion exchange chromatography  
 Japanese encephalitis virus  
 Lassa virus  
 Measles virus  
 Mumps virus  
 Orthomyxovirus  
 Paramyxovirus  
 Phenyl group  
 Poxviridae  
 Purification  
 Rabies virus  
 Reoviridae  
 Respiratory syncytial virus

Retroviridae  
 Rubella virus  
 Russian spring summer encephalitis virus  
 SARS coronavirus  
 Sendai virus  
 Size-exclusion chromatography  
 Surfactants  
 Temperature  
 Togaviridae  
 Variola virus  
 Vesicular stomatitis virus  
 Virus  
 West Nile virus  
 Yellow fever virus

pH

(method for purifying virus envelope by column chromatog.)

IT Biopolymers

Gene

RL: BCP (Biochemical process); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(method for purifying virus envelope by column chromatog.)

IT Functional groups

(oligoethyleneglycol; method for purifying virus envelope by column chromatog.)

IT Solvents

(organic, hydrophilic; method for purifying virus envelope by column chromatog.)

IT Alcohols, uses

RL: TEM (Technical or engineered material use); USES (Uses)

(polyhydric; method for purifying virus envelope by column chromatog.)

IT Anion exchange chromatography

(weakly basic, diethylaminopropyl (DEAP); method for purifying virus envelope by column chromatog.)

IT 9001-67-6, Neuraminidase

RL: ANT (Analyte); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)

(method for purifying virus envelope by column chromatog.)

IT 107-21-1, Ethyleneglycol, uses 7439-95-4, Magnesium, uses 7440-70-2, Calcium, uses 7447-40-7, Potassium chloride, uses 7647-14-5, Sodium chloride, uses 7757-82-6, Sodium sulfate, uses 7783-20-2, Ammonium sulfate, uses 9002-93-1, Triton X-100 9005-65-6, Tween 80 9012-36-6, Sepharose

RL: TEM (Technical or engineered material use); USES (Uses)

(method for purifying virus envelope by column chromatog.)

RE.CNT 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

- (1) Alain, J; Protein Expression and Purification 1995, V6, P91
- (2) Anges Mg Inc; EP---1420065 A1 2003 HCAPLUS
- (3) Anges Mg Inc; AU2002318581 A1 2003
- (4) Anges Mg Inc; WO2003014338 A1 2003
- (5) Anges Mg Inc; JP2003519468 X 2003
- (6) Anges Mg Inc; US2004253272 A1 2003 HCAPLUS
- (7) Avant Immunotherapeutics Inc; AU---9896956 A 1999 HCAPLUS
- (8) Avant Immunotherapeutics Inc; WO---9919345 A1 1999 HCAPLUS
- (9) Chiron Corp; JP-05-505616 A 1993
- (10) Chiron Corp; EP----519001 A1 1993 HCAPLUS
- (11) Chiron Corp; DE--69132795 E 1993
- (12) Chiron Corp; IE-----83584 B 1993
- (13) Chiron Corp; WO---9113906 A 1993 HCAPLUS
- (14) Chiron Corp; PT-----96994 A 1993
- (15) Marcus, S; Virology 1978, V86(2), P398 HCAPLUS
- (16) Teramoto, Y; Journal of Virology 1979, V31(2), P334 HCAPLUS
- (17) Welling, G; J Chromatogr 1984, V297, P101 HCAPLUS
- (18) Yamanouchi Pharm Co Ltd; JP-03-505322 X 1991
- (19) Yamanouchi Pharm Co Ltd; WO---9113976 A 1991 HCAPLUS

L19 ANSWER 2 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN  
 AN 2005:394045 HCAPLUS  
 DN 142:426441  
 ED Entered STN: 09 May 2005  
 TI Enzyme activities and pH tests for the determination of the risk of  
 obstetric and gynecologic complications in samples of body fluids of women  
 IN Cauci, Sabina  
 PA Unibio S.R.L., Italy  
 SO Eur. Pat. Appl., 19 pp.  
 CODEN: EPXXDW  
 DT Patent  
 LA English  
 IC ICM G01N-0033/569  
 ICS G01N-0033/50; C12Q-0001/34; C12Q-0001/37; G01N-0033/48  
 CC 9-16 (Biochemical Methods)  
 Section cross-reference(s): 14

## FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP---1528396	A1	20050504	2004EP-0022918	20040927 <--
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR				
	US2005095660	A1	20050505	2003US-0698795	20031031 <--
	CA---2485854	AA	20050430	2004CA-2485854	20041025 <--
PRAI	2003US-0698795	A	20031031	<--	

## CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
EP 1528396	ICM	G01N-0033/569
	ICS	G01N-0033/50; C12Q-0001/34; C12Q-0001/37; G01N-0033/48
	IPCI	G01N0033-569 [ICM,7]; G01N0033-50 [ICS,7]; C12Q0001-34 [ICS,7]; C12Q0001-37 [ICS,7]; G01N0033-48 [ICS,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]; C12Q0001-37 [I,A]; C12Q0001-37 [I,C]; G01N0033-68 [I,A]; G01N0033-68 [I,C]
US2005095660	ECLA	C12Q001/34; C12Q001/37; G01N033/68V <--
	IPCI	C12Q0001-34 [ICM,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]; C12Q0001-37 [I,A]; C12Q0001-37 [I,C]; G01N0033-68 [I,A]; G01N0033-68 [I,C]
	NCL	435/018.000
CA---2485854	ECLA	C12Q001/34; C12Q001/37; G01N033/68V <--
	IPCI	C12Q0001-37 [ICM,7]; C12Q0001-34 [ICS,7]; G01N0033-52 [ICS,7]; G06F0017-60 [ICS,7]; G01N0033-84 [ICS,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]; C12Q0001-37 [I,A]; C12Q0001-37 [I,C]; G01N0033-68 [I,A]; G01N0033-68 [I,C]
	ECLA	C12Q001/34; C12Q001/37; G01N033/68V <--

AB The current invention describes a method for selecting a particular population of women having a risk of developing obstetric or gynecol. pathologies indicated as odds ratio (OR) value higher than 5.5, comprising the following steps in order: (a) determination of the levels of **sialidase** by means of the procedure described in Cauci et al. Am J Obstet Gynecol. 1998; 178; 511-5 and/or **prolidase** activity by means of the procedure described in Cauci et al. J Infect Dis 1998; 178; 1698-706 in samples of body fluid; (b) determination of the pH value of said body fluid samples; (c) selecting the samples having a **sialidase** value equal or above 5.0 nmol of methoxyphenol and/or a **prolidase** level equal or above 1500 MOD for **prolidase** and a pH  $\geq$  5.0. Consequently, this method gives the physician an efficient tool to decide whether or not to administer a pharmacol. therapy to women at risk of severe adverse outcomes.

ST enzyme activity pH test detn risk obstetric gynecol

IT Body fluid  
 Computer program  
 Human

## Test kits

## pH

(enzyme activities and pH tests for determination of risk of obstetric and gynecol. complications in samples of body fluids of women)

IT Medicine

(gynecol.; enzyme activities and pH tests for determination of risk of obstetric and gynecol. complications in samples of body fluids of women)

IT Medicine

(obstetrics; enzyme activities and pH tests for determination of risk of obstetric and gynecol. complications in samples of body fluids of women)

IT 9001-67-6, Sialidase 9025-32-5,

## Prolidase

RL: ANT (Analyte); BSU (Biological study, unclassified); ANST

(Analytical study); BIOL (Biological study)

(enzyme activities and pH tests for determination of risk of obstetric and gynecol. complications in samples of body fluids of women)

IT 3304-59-4 3326-64-5 7369-91-7, L-Proline-p-nitroanilide 16037-15-3,

L-Proline- $\beta$ -naphthylamide 24751-40-4 26112-88-9 76204-02-9

86925-99-7 94720-65-7 96643-94-6 153248-52-3

RL: ARU (Analytical role, unclassified); ANST (Analytical study)

(enzyme activities and pH tests for determination of risk of obstetric and gynecol. complications in samples of body fluids of women)

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD

## RE

(1) Cauci, S; WO--02065122 A 2002 HCAPLUS

(2) Cauci, S; WO--02065130 A 2002 HCAPLUS

(3) Cauci, S; JOURNAL OF CLINICAL MICROBIOLOGY 2003, V41(1), P435 HCAPLUS

(4) Cauci, S; JOURNAL OF INFECTIOUS DISEASES 1998, V178(6), P1698 HCAPLUS

L19 ANSWER 3 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2002:637929 HCAPLUS

DN 137:167678

ED Entered STN: 23 Aug 2002

TI Enzymatic test for the determination of the risk of pathologies related to the presence of sialidase or prolidase activity in women body fluid samples

IN Cauci, Sabina

PA Unibio S.R.L., Italy

SO PCT Int. Appl., 27 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM G01N-0033/50

ICS C12Q-0001/34; C12Q-0001/37

CC 14-13 (Mammalian Pathological Biochemistry)

Section cross-reference(s): 1, 7

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO2002065122	A1	20020822	2001WO-IT00069	20010215
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
EP---1360484	A1	20031112	2001EP-0912101	20010215
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR			
CN---1503908	A	20040609	2001CN-0822652	20010215
US2004219617	A1	20041104	2003US-0467357	20031020

PRAI 2001WO-IT00069

W

20010215

## CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2002065122	ICM	G01N-0033/50
	ICS	C12Q-0001/34; C12Q-0001/37
	IPCI	G01N0033-50 [ICM,7]; C12Q0001-34 [ICS,7]; C12Q0001-37 [ICS,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]; C12Q0001-37 [I,A]; C12Q0001-37 [I,C]; G01N0033-50 [I,A]; G01N0033-50 [I,C]
	ECLA	C12Q001/34; C12Q001/37; G01N033/50D4
EP---1360484	IPCI	G01N0033-50 [ICM,7]; C12Q0001-34 [ICS,7]; C12Q0001-37 [ICS,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]; C12Q0001-37 [I,A]; C12Q0001-37 [I,C]; G01N0033-50 [I,A]; G01N0033-50 [I,C]
CN---1503908	IPCI	G01N0033-50 [ICM,7]; C12Q0001-34 [ICS,7]; C12Q0001-37 [ICS,7]
US2004219617	IPCI	G01N0033-554 [ICM,7]; G01N0033-569 [ICS,7]; C12Q0001-26 [ICS,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]; C12Q0001-37 [I,A]; C12Q0001-37 [I,C]; G01N0033-50 [I,A]; G01N0033-50 [I,C]
	NCL	435/007.320
	ECLA	C12Q001/34; C12Q001/37; G01N033/50D4
AB	The current invention describes a method for the determination of the risk of pathologies related to the presence of sialidase and/or prolidase activity in body fluid samples of women, comprising the following steps in order: (a) determination of the levels of sialidase and/or prolidase activity in said sample of body fluid; (b) comparison of said levels of sialidase and/or prolidase activity with ranges of prefixed values of said activity; (c) calcn. of the risk factor. This method was particularly efficient in permitting an accurate and reliable evaluation of the risk of pathologies related to the presence of sialidase and/or prolidase activity in samples of body fluid of women. Consequently, this method gives the physician an efficient tool to decide whether or not to administer a pharmacol. therapy.	
ST	sialidase prolidase detn body fluid risk pathol; pregnancy sialidase prolidase body fluid	
IT	Vagina (anal. of fluid of; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)	
IT	Infection (bacterial, vaginosis; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)	
IT	Vagina, disease (bacterial; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)	
IT	Inflammation Uterus, disease (cervicitis; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)	
IT	Inflammation Uterus, disease (endometritis; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)	
IT	Acid-base indicators Body fluid Diagnosis	

Disease, animal  
Gardnerella vaginalis  
Human

Pregnancy  
Risk assessment

Test kits

(enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)

- IT Fertility disorders  
(female, from upper genital tract infections; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Pregnancy  
(first trimester; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Fluorescent substances  
(fluorogenic substrates; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Surgery  
(gynecol., infection after; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Uterus, disease  
(infection, post-partum; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Reproductive system, disease  
(infection, upper; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Amniotic fluid  
(intraamniotic infection; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Parturition  
(low weight at; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Body weight  
(low, at birth; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT pH  
(of vaginal fluid sample; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Amnion, disease  
(premature rupture; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Parturition  
(premature; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Infection  
(reproductive system, upper; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Pregnancy  
(second trimester; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Abortion



- (spontaneous; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Color formers  
(substrates; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Human immunodeficiency virus  
(susceptibility to sexually or vertically transmitted infection with; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Human papillomavirus  
Papillomavirus  
(susceptibility to sexually transmitted infection with; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Sexually transmitted diseases  
(susceptibility to; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT Infection  
(uterine, post-partum; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT 9001-67-6, Sialidase 9025-32-5,  
Prolidase  
RL: ANT (Analyte); BSU (Biological study, unclassified); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
(enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT 3304-59-4, N-Benzoyloxycarbonyl-L-proline-p-nitrophenyl ester 3326-64-5  
7369-91-7, L-Proline-p-nitroanilide 16037-15-3, L-Proline- $\beta$ -naphthylamide 86925-99-7 94720-65-7 96643-94-6  
RL: ARG (Analytical reagent use); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
(prolidase reagent; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)
- IT 24751-40-4 76204-02-9 153248-52-3 157707-92-1  
RL: ARG (Analytical reagent use); DGN (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES (Uses)  
(sialidase reagent; enzymic test for determination of risk of pathol. related to presence of sialidase or prolidase activity in women body fluid samples)

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

- (1) Briselden, A; JOURNAL OF CLINICAL MICROBIOLOGY 1992, V30(3), P663 HCAPLUS
- (2) Corfield, T; WO---0055354 A 2000 HCAPLUS
- (3) Ibbex Inc; WO---0024753 A 2000 HCAPLUS
- (4) Lawrence, P; US---5571684 A 1996 HCAPLUS
- (5) McGregor, J; AMERICAN JOURNAL OF OBSTETRICS & GYNECOLOGY 1994, V170(4), P1048 MEDLINE

L19 ANSWER 4 OF 4 HCAPLUS COPYRIGHT 2006 ACS on STN

AN 2000:725795 HCAPLUS

DN 133:263206

ED Entered STN: 13 Oct 2000

TI Method for detecting and assaying exoglycosidase activity

IN Zhu, Alex

PA New York Blood Center, Inc., USA

SO PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DT Patent

LA English  
 IC ICM C12Q-0001/34  
 ICS C12Q-0001/54; C12Q-0001/00; C12Q-0001/37; G01N-0033/53  
 CC 7-1 (Enzymes)  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO2000060111	A1	20001012	2000WO-US09053	20000405
	W: CA, JP, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	US---6171810	B1	20010109	1999US-0287869	19990407
PRAI	1999US-0287869	A	19990407		

## CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2000060111	ICM	C12Q-0001/34
	ICS	C12Q-0001/54; C12Q-0001/00; C12Q-0001/37; G01N-0033/53
	IPCI	C12Q0001-34 [ICM,7]; C12Q0001-54 [ICS,7]; C12Q0001-00 [ICS,7]; C12Q0001-37 [ICS,7]; G01N0033-53 [ICS,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]
	ECLA	C12Q001/34
US---6171810	IPCI	C12Q0001-34 [ICM,7]; C12Q0001-54 [ICS,7]; C12Q0001-00 [ICS,7]
	IPCR	C12Q0001-34 [I,A]; C12Q0001-34 [I,C]
	NCL	435/018.000; 435/004.000; 435/014.000; 435/968.000; 536/001.110; 536/123.100; 536/123.130
	ECLA	C12Q001/34

AB A method for detecting and measuring exoglycosidase activity is presented. The method employs derivs. containing the fluorescent group 4-methylumbelliferyl ("4-Mu") at a pH lower than that conventionally employed. While the fluorescence intensity due to the 4-Mu group is considerably diminished at the lower pHs employed, the fluorescent intensity is still sufficient to continuously measure exoglycosidase activity in the activity range commonly assayed. The method is easily adaptable to high throughput enzyme assay systems and automated data anal. method. The method also provides a means to detect alterations in exoglycosidase activity that are independent of expression levels. The figure shows the pH dependence of 4-Mu fluorescence intensity over a pH range between 3 and 10, when measured with an excitation wavelength of 365 nm and an emission wavelength of 440 nm, and at concns. of 1 and 10 nM, wherein (O) corresponds to 1 nM 4-Mu, and (Δ) corresponds to 10 nM 4-Mu.

ST detecting assaying exoglycosidase activity

IT Functional groups

(4-methylumbelliferyl; method for detecting and assaying exoglycosidase activity)

IT Fluorometry

pH

(method for detecting and assaying exoglycosidase activity)

IT Enzymes, biological studies

RL: ANT (Analyte); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)

(method for detecting and assaying exoglycosidase activity)

IT 9001-67-6, Neuraminidase 9025-35-8,  
 α-Galactosidase 9037-65-4, α-Fucosidase 9075-63-2,  
 α-N-Acetylgalactosaminidase 52769-51-4, Endoglycosidase  
 52769-52-5, Exoglycosidase

RL: ANT (Analyte); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)

(method for detecting and assaying exoglycosidase activity)

IT 38597-12-5, 4-Methylumbelliferyl-α-D-galactoside 54322-38-2,  
 4-Methylumbelliferyl-α-L-fucoside 59322-44-0, 4-Methylumbelliferyl-N-acetyl-α-D-neuraminic acid

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)  
 (method for detecting and assaying exoglycosidase activity)

RE.CNT 2        THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD  
RE  
(1) Miles; US---3850322 A 1974  
(2) Robinson; Clinica Chimica Acta V55, P65 HCAPLUS

=> => b medl  
FILE 'MEDLINE' ENTERED AT 17:15:44 ON 02 MAR 2006

FILE LAST UPDATED: 1 MAR 2006 (20060301/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 is now (26 Feb.) available. For details  
on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>).  
See also:

<http://www.nlm.nih.gov/mesh/>  
[http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\\_mesh.html](http://www.nlm.nih.gov/pubs/techbull/nd04/nd04_mesh.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_med\\_data\\_changes.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_med_data_changes.html)  
[http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\\_2006\\_MeSH.html](http://www.nlm.nih.gov/pubs/techbull/nd05/nd05_2006_MeSH.html)

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the  
MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate  
substance identification.

=> d all 128 tot

L28 ANSWER 1 OF 4        MEDLINE on STN  
AN 97436569        MEDLINE  
DN PubMed ID: 9292542  
TI Presence in human erythrocyte membranes of a novel form of  
sialidase acting optimally at neutral pH.  
AU Venerando B; Fiorilli A; Croci G L; Tettamanti G  
CS Department of Medical Chemistry and Biochemistry, The Medical School,  
University of Milan, Italy.  
SO Blood, (1997 Sep 1) Vol. 90, No. 5, pp. 2047-56.  
Journal code: 7603509. ISSN: 0006-4971.  
CM Comment in: Blood. 2002 Aug 15;100(4):1511. PubMed ID: 12184275  
CY United States  
DT Journal; Article; (JOURNAL ARTICLE)  
LA English  
FS Abridged Index Medicus Journals; Priority Journals  
EM 199709  
ED Entered STN: 19971013  
Last Updated on STN: 19971013  
Entered Medline: 19970930  
AB The feature of intact human erythrocytes and erythrocyte white ghosts is a  
unique sialidase activity with acidic optimal pH (acidic  
sialidase). The treatment of white ghosts with mildly alkaline  
isotonic solutions at 37 degrees C, like that used to produce resealed  
ghosts, is accompanied by the expression, together with the acidic  
sialidase, of a novel sialidase with a pH optimum of 7.2  
(neutral sialidase) that remained masked in the inside-out  
vesicles prepared from white ghosts. Exhaustive treatment of resealed  
ghosts with Bacillus Thuringiensis phosphatidylinositol-specific  
phospholipase C causes an almost complete release of the acidic  
sialidase, with the neutral enzyme remaining totally unaffected.  
The treatment of resealed ghosts with 1.2% Triton X-100 resulted in the  
solubilization of only the neutral sialidase, whereas 3.6%  
octylglucoside also solubilized the acidic sialidase. The  
neutral enzyme affected not only the artificial substrate but also any

sialoderivatives of a ganglioside, glycoprotein, and oligosaccharide nature; the acidic enzyme did not affect sialoglycoproteins. Erythrocyte endogenous gangliosides were hydrolyzed by both sialidases, whereas the endogenous sialoglycoproteins responded to only the neutral enzyme. It was definitely proved that the acidic sialidase is located on the outer erythrocyte membrane surface, so presumably the neutral enzyme has the same location. It could be that the newly discovered neutral sialidase has a physiologic role in the releasing of sialic acid from erythrocytes during the erythrocyte aging process, leading to eventual phagocytosis by macrophages.

CT Cell Aging

\*Erythrocyte Membrane: EN, enzymology

Humans

Hydrogen-Ion Concentration

\*Neuraminidase: AN, analysis

Neuraminidase: CH, chemistry

Neuraminidase: ME, metabolism

Research Support, Non-U.S. Gov't

CN EC 3.2.1.18 (Neuraminidase)

L28 ANSWER 2 OF 4 MEDLINE on STN

AN 75191973 MEDLINE

DN PubMed ID: 238036

TI Preparation of a glycoprotein fraction from pooled human plasma and its evaluation as a substrate for the assay of *Clostridium welchii* (*C. perfringens*) neuraminidase.

AU Fraser A G; Smith J K

SO Journal of medical microbiology, (1975 May) Vol. 8, No. 2, pp. 235-49.

Journal code: 0224131. ISSN: 0022-2615.

CY ENGLAND: United Kingdom

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

EM 197509

ED Entered STN: 19900310

Last Updated on STN: 19970203

Entered Medline: 19750924

AB A glycoprotein fraction (fraction VII) suitable for use as a substrate in assays of microbial neuraminidase was prepared from pooled human plasma. It is pasteurised during preparation to eliminate the risk of transmission of serum hepatitis. This results in polymerisation of some of the gamma1-acid glycoprotein, but fraction VII is shown to be an excellent substrate for the neuraminidase of *Clostridium welchii* (*C. perfringens*). A sensitive assay procedure is described. A number of factors may interfere with the measurement of NANA released by the action of microbial neuraminidase and procedures are described for evaluation of this problem. Fraction VII is easy to prepare, cheap and available in standard form in large amounts (inquiries should be addressed to J. K. S.); it is recommended for routine use as a convenient substrate for neuraminidase assays.

CT \*Clostridium perfringens: EN, enzymology

Culture Media

Dialysis

Dose-Response Relationship, Drug

Glycoproteins: AN, analysis

\*Glycoproteins: BL, blood

Glycoproteins: ME, metabolism

Humans

Hydrogen-Ion Concentration

Kinetics

\*Neuraminidase: AN, analysis

Neuraminidase: ME, metabolism

Sialic Acids: ME, metabolism

CN 0 (Culture Media); 0 (Glycoproteins); 0 (Sialic Acids); EC 3.2.1.18 (Neuraminidase)

L28 ANSWER 3 OF 4 MEDLINE on STN  
 AN 74306651 MEDLINE  
 DN PubMed ID: 4137161  
 TI Red cell hydrolases. 3. Neuraminidase activity in isolated human erythrocyte plasma membranes.  
 AU Bosmann H B  
 SO Vox sanguinis, (1974) Vol. 26, No. 6, pp. 497-512.  
 Journal code: 0413606. ISSN: 0042-9007.  
 CY Switzerland  
 DT Journal; Article; (JOURNAL ARTICLE)  
 LA English  
 FS Priority Journals  
 EM 197410  
 ED Entered STN: 19900310  
 Last Updated on STN: 19900310  
 Entered Medline: 19741017  
 CT Check Tags: Male  
 Alpha-Globulins: ME, metabolism  
 Blood Protein Electrophoresis  
 Blood Proteins: AN, analysis  
 Borohydrides: ME, metabolism  
 Cell Membrane: EN, enzymology  
 Electrophoresis, Polyacrylamide Gel  
 Erythrocytes: DE, drug effects  
 \*Erythrocytes: EN, enzymology  
 Fetal Proteins: ME, metabolism  
 Humans  
 Hydrogen-Ion Concentration  
 \*Neuraminidase: ME, metabolism  
 Neuraminidase: PD, pharmacology  
 Potassium  
 Sodium Dodecyl Sulfate  
 Surface-Active Agents  
 Temperature  
 Tritium  
 RN 10028-17-8 (Tritium); 151-21-3 (Sodium Dodecyl Sulfate); 7440-09-7 (Potassium)  
 CN 0 (Alpha-Globulins); 0 (Blood Proteins); 0 (Borohydrides); 0 (Fetal Proteins); 0 (Surface-Active Agents); EC 3.2.1.18 (Neuraminidase)

L28 ANSWER 4 OF 4 MEDLINE on STN  
 AN 72066676 MEDLINE  
 DN PubMed ID: 5128828  
 TI Neuraminidase activity in human leukocytes.  
 AU Yeh A K; Tulsiani D R; Carubelli R  
 SO The Journal of laboratory and clinical medicine, (1971 Nov) Vol. 78, No. 5, pp. 771-8.  
 Journal code: 0375375. ISSN: 0022-2143.  
 CY United States  
 DT Journal; Article; (JOURNAL ARTICLE)  
 LA English  
 FS Abridged Index Medicus Journals; Priority Journals  
 EM 197202  
 ED Entered STN: 19900310  
 Last Updated on STN: 19970203  
 Entered Medline: 19720223  
 CT Bile Acids and Salts  
 Calcium: PD, pharmacology  
 Chlorides: PD, pharmacology  
 Copper: PD, pharmacology  
 Freezing  
 Humans  
 Hydrogen-Ion Concentration  
 Hydrolysis

Ions

Kinetics

Leukocytes: DE, drug effects

\*Leukocytes: EN, enzymology

Magnesium: PD, pharmacology

Mercury: PD, pharmacology

\*Neuraminidase: AN, analysis

Refrigeration

Surface-Active Agents: PD, pharmacology

Zinc: PD, pharmacology

RN 7439-95-4 (Magnesium); 7439-97-6 (Mercury); 7440-50-8 (Copper); 7440-66-6 (Zinc); 7440-70-2 (Calcium)

CN 0 (Bile Acids and Salts); 0 (Chlorides); 0 (Ions); 0 (Surface-Active Agents); EC 3.2.1.18 (Neuraminidase)

=&gt; =&gt; d his

(FILE 'HOME' ENTERED AT 16:43:23 ON 02 MAR 2006)

FILE 'HCAPLUS' ENTERED AT 16:43:35 ON 02 MAR 2006

L1 1 US2005095660/PN OR US2003-698795#/AP, PRN  
E CAUCI S/AU  
L2 25 E3-4  
L3 3 UNIBIO/CS, PA

FILE 'REGISTRY' ENTERED AT 16:45:29 ON 02 MAR 2006

FILE 'HCAPLUS' ENTERED AT 16:45:32 ON 02 MAR 2006

L4 TRA L1 1- RN : 13 TERMS

FILE 'REGISTRY' ENTERED AT 16:45:32 ON 02 MAR 2006

L5 13 SEA L4  
SEL RN 9-10  
L6 2 E1-2 AND L5

FILE 'HCAPLUS' ENTERED AT 16:50:16 ON 02 MAR 2006

L7 5992 L6  
L8 549 DIPEPTIDASE (1A) PROLINE OR PROLIDASE OR E C ( ) (3 4 13 9 OR 3 4  
L9 13682 NEURAMINIDASE OR ACETYLNEURAMINIDASE? OR ARYLNEURAMINIDASE? OR  
L10 244 L7-9 (L) ANT/RL  
L11 2 L10 AND L1-3  
L12 242 L10 NOT L11  
E PH/CT  
L13 36489 E3-4  
E E3+ALL  
L14 50840 E7+OLD, NT  
L15 2 L12 AND L13-14  
L16 4 L11, L15  
L17 2 L16 AND L1-3  
L18 4 L16 AND L7-15  
L19 4 L17-18

FILE 'MEDLINE' ENTERED AT 17:01:09 ON 02 MAR 2006

L20 15221 L7-9  
E PH/CT  
E E3+ALL  
E E2+ALL  
L21 197732 E5+NT  
L22 1009 L20 AND L21  
E BODY FLUID/CT  
E E3+ALL  
E E2+ALL  
L23 270 E3+NT AND L22  
L24 268 L23 AND PY<=2003  
E NEURAMINIDASE/CT

L25           E E3+ALL  
           554 E6 (L)AN/CT  
           E SIALIDASE/CT  
           E E3+ALL  
           E PROLIDASE/CT  
 L26           4588 L20 AND AN/CT  
 L27           83 L24 AND L25-26  
           SEL AN 4 48 55 71  
 L28           4 L27 AND E1-4

FILE 'EMBASE' ENTERED AT 17:15:57 ON 02 MAR 2006  
 L29           12036 L7-9

          E SIALIDASE/CT  
           E E3+ALL

L30           5690 E1  
           E PROLIDASE/CT  
           E E3+ALL  
           E E2+ALL

L31           309 E1  
           E PH/CT  
           E E3+ALL

L32           159 E5+NT AND L29-31  
           E BODY FLUID/CT  
           E E3+ALL

L33           28 E3+NT AND L32

FILE 'HCAPLUS' ENTERED AT 17:29:12 ON 02 MAR 2006  
 L34           14222 L8-9

          SAV TEM L34 GIT795F0/A

FILE 'REGISTRY' ENTERED AT 17:30:34 ON 02 MAR 2006  
           SAV TEM GIT795F1/A L6

=> b embase

FILE 'EMBASE' ENTERED AT 17:59:33 ON 02 MAR 2006  
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FILE COVERS 1974 TO 24 Feb 2006 (20060224/ED)

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This file contains CAS Registry Numbers for easy and accurate  
 substance identification.

=> d all 140 tot

L40 ANSWER 1 OF 1 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights  
       reserved on STN  
 AN    2002161524 EMBASE  
 TI    Different behavior of ghost-linked acidic and neutral sialidases  
       during human erythrocyte ageing.  
 AU    Tringali C.; Fiorilli A.; Venerando B.; Tettamanti G.  
 CS    Prof. G. Tettamanti, Department of Medical Chemistry, Medical School,  
       University of Milan, via Fratelli Cervi 93, 20090 Segrate (Milan), Italy.  
       guido.tettamanti@unimi.it  
 SO    Glycoconjugate Journal, (2001) Vol. 18, No. 5, pp. 407-418.  
       Refs: 65  
       ISSN: 0282-0080 CODEN: GLJOEW  
 CY    Netherlands  
 DT    Journal; Article  
 FS    025 Hematology  
       029 Clinical Biochemistry  
 LA    English  
 SL    English  
 ED    Entered STN: 20020523  
       Last Updated on STN: 20020523

AB Acidic and neutral sialidases (pH optimum 4.7 and 7.2, respectively) were assayed on human circulating erythrocytes during ageing. The assays were performed on intact erythrocytes and resealed erythrocyte ghost membranes. From young to senescent erythrocytes the acidic sialidase featured a 2.7-fold and 2.5-fold decrease in specific activity when measured on intact cells or resealed ghost membranes, whereas the neutral sialidase a 5-fold and 7-fold increase, respectively. The Ca(2+)-loading procedure was employed to mimic the vesiculation process occurring during erythrocyte ageing. Under these conditions the released vesicles displayed an elevated content of acidic sialidase, almost completely linked through a glycan phosphoinositide (GPI) anchor but no neutral sialidase activity, that was completely retained by remnant erythrocytes together with almost all the starting content of sialoglycoconjugates. The loss with vesiculation of acidic sialidase with a concomitant relative increase of neutral sialidase was more marked in young than senescent erythrocytes. The data presented suggest that during ageing erythrocytes loose acidic sialidase, and get enriched in the neutral enzyme, the vesiculation process, possibly involving GPI-anchors-rich membrane microdomains, being likely responsible for these changes. The enhanced neutral sialidase activity might account for the sialic acid loss occurring during erythrocyte ageing.

## CT Medical Descriptors:

\*erythrocyte lifespan

\*erythrocyte ghost

enzyme activity

pH

membrane vesicle

erythrocyte membrane

density gradient centrifugation

human

male

female

controlled study

human cell

adult

article

priority journal

## Drug Descriptors:

\*sialidase: EC, endogenous compound

glycan

phosphatidylinositol

glycoconjugate

calcium ion

ganglioside

sialic acid

RN (sialidase) 9001-67-6; (calcium ion) 14127-61-8

=&gt; b biosis

FILE 'BIOSIS' ENTERED AT 17:59:40 ON 02 MAR 2006

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FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT  
FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 1 March 2006 (20060301/ED)

=&gt; d all 138 tot

L38 ANSWER 1 OF 1 BIOSIS. COPYRIGHT (c) 2006 The Thomson Corporation on STN

AN 2005:159180 BIOSIS

DN PREV200500166039

TI Combination of vaginal pH with vaginal sialidase and



prolidase activities for prediction of low birth weight and preterm birth.

AU Cauci, Sabina [Reprint Author]; McGregor, James; Thorsen, Poul;  
Grove, Jakob; Guaschino, Secondo

CS Dipartimento Sci and Tecnol Biomed, Fac Med and Chirurg, Piazzale Kolbe 4,  
I-33100, Udine, Italy  
scauci@mail.dstb.uniud.it

SO American Journal of Obstetrics and Gynecology, (February 2005) Vol. 192,  
No. 2, pp. 489-496, 478. print.  
CODEN: AJOGAH. ISSN: 0002-9378.

DT Article

LA English

ED Entered STN: 27 Apr 2005  
Last Updated on STN: 27 Apr 2005

AB Objective: The purpose of this study was to assess if easy to measure vaginal fluid biomarkers are predictive for low birth weight (LBW, < 2500 g), very LBW (VLBW, <1500 g), spontaneous preterm at <37 weeks' gestation, and total preterm, deliveries (at <37, <35, <32 weeks' gestation). Study design: Low and high cutoffs for vaginal fluid pH, sialidase, and prolidase activities were examined in a nested case-control study of 579 Danish women (from a study population of 2846 women) with samples collected at mean 17 weeks' gestation. One hundred sixteen LBW (17 VLBW), 117 preterm deliveries (85 spontaneous), and 418 normal term deliveries were analyzed. Results: Vaginal pH gtoreq4.7 or pH gtoreq5 by itself was not associated with LBW or prematurity. Conversely, combination of pH 5 and high sialidase activity demonstrated OR 17 (CI 1.8150) for LBW OR 31 (CI 1.8-516) for VLBW; along with OR 18 (CI 1.6-204) for preterm at <35 weeks'; and OR 31 (CI 1.9-542) for preterm at <32 weeks' gestation. The combination of pH gtoreq5 and high prolidase activity demonstrated OR 13 (CI 1.3-122) for LBW; OR 33 (CI 2.0-553) for VLBW, as well as OR 9.2 (CI 6.6-150) for preterm at <35 weeks'; and OR 35 (CI 2.0-586) for preterm at <32 weeks' gestation. In this population, no woman having high sialidase and high prolidase activity had a term birth, or a baby weighting 2500 g at birth. Conclusion: In this Danish population, inid-gestation findings of vaginal fluid elevated pH with sialidase and/or prolidase were associated with LBW, VLBW, and early preterm at <35 or <32 weeks' gestation. Copyright 2005 Elsevier Inc. All rights reserved.

CC Clinical biochemistry - General methods and applications 10006  
Enzymes - General and comparative studies: coenzymes 10802  
Reproductive system - Physiology and biochemistry 16504  
Reproductive system - Pathology 16506  
Pediatrics 25000

IT Major Concepts  
Clinical Chemistry (Allied Medical Sciences); Gynecology (Human Medicine, Medical Sciences); Obstetrics (Human Medicine, Medical Sciences)

IT Parts, Structures, & Systems of Organisms  
vaginal fluid: reproductive system

IT Diseases  
preterm birth: reproductive system disease/female Labor, Premature (MeSH)

IT Chemicals & Biochemicals  
prolidase [EC 3.4.13.9]; sialidase

IT Miscellaneous Descriptors  
low birth weight; vaginal pH

ORGN Classifier  
Hominidae 86215  
Super Taxa  
Primates; Mammalia; Vertebrata; Chordata; Animalia  
Organism Name  
human (common): adolescent, adult, Danish, female  
Taxa Notes  
Animals, Chordates, Humans, Mammals, Primates, Vertebrates

RN 9025-32-5 (prolidase)  
9025-32-5 (EC 3.4.13.9)  
9001-67-6 (sialidase)

=> d his l35-

FILE 'BIOSIS' ENTERED AT 17:56:48 ON 02 MAR 2006

L35 14273 L7-9  
E CAUCI S/AU  
L36 36 E3-4  
L37 11 L35 AND L36  
L38 1 L37 AND (PH OR HYDROGEN (1W) ION)  
L39 1061 L35 AND (PH OR HYDROGEN (1W) ION)

FILE 'EMBASE' ENTERED AT 17:59:15 ON 02 MAR 2006

SEL AN 13 L33  
L40 1 E1 AND L33

=> b biosis

FILE 'BIOSIS' ENTERED AT 08:17:26 ON 03 MAR 2006

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FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNS) PRESENT  
FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 1 March 2006 (20060301/ED)

=> d all 116 tot

L16 ANSWER 1 OF 1 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

AN 1985:238953 BIOSIS

DN PREV198579018949; BA79:18949

TI STUDIES ON PRENATAL DIAGNOSIS OF HEREDITARY LYSOSOMAL STORAGE DISEASES.

AU WAGATSUMA K [Reprint author]

CS DEP PEDIATR, SAPPORO MED COLL, JPN

SO Sapporo Medical Journal, (1984) Vol. 53, No. 4, pp. 373-394.

CODEN: SIZSAR. ISSN: 0036-472X.

DT Article

FS BA

LA JAPANESE

AB Assay conditions were studied for 11 lysosomal enzymes ( $\beta$ -D-galactosidase,  $\alpha$ -D-mannosidase,  $\beta$ -hexosaminidase,  $\beta$ -D-glucuronidase,  $\alpha$ -D-galactosidase,  $\alpha$ -D-glucosidase, arylsulfatase,  $\beta$ -D-glucosidase,  $\alpha$ -L-fucosidase,  $\alpha$ -D-neuraminidase and  $\alpha$ -L-iduronidase) in cultured amniotic fluid cells(CAFC), cultured skin fibroblasts(CSF) and cultured embryonic lung fibroblasts(CELF), and the specific activities of the enzymes were compared among these cultured cells. In addition, changes in these enzymes from the 3 cell types were investigated between 4-6 earlier passages and 24-26 later passages, with regard to their specific activities, Km values and pH profiles. The following results were obtained. All enzymes assayed for the 4-6 earlier passages had the same Km values for CAFC, CSF and CELF. With the exception of  $\alpha$ -D-neuraminidase and  $\alpha$ -L-fucosidase, the enzymes also had the same pH optima. The specific activities of  $\beta$ -D-glucuronidase, arylsulfatase,  $\alpha$ -D-glucosidase and  $\beta$ -D-glucosidase significantly increased with development. All enzymes assayed in the 3 cell types were also unchanged with cell aging, with regard to their Km values. With the exception of  $\alpha$ -D-glucosidase,  $\alpha$ -D-neuraminidase and  $\alpha$ -L-fucosidase, the enzymes were also unchanged in their points of pH optima. No changes were observed with development in the specific activities of  $\beta$ -D-glucosidase,  $\beta$ -D-glucuronidase,  $\alpha$ -D-galactosidase,  $\alpha$ -D-mannosidase,  $\beta$ -D-galactosidase,  $\beta$ -hexosaminidase and  $\alpha$ -D-neuraminidase from the 3 cell types. Variations were observed between the levels of these enzymes in the 3 cell types with cell aging, such as increases in some, decreases in others and no change in still others. Especially, the specific activities of  $\alpha$ -D-mannosidase in CAFC and CSF and those of  $\alpha$ -L-fucosidase in CELF markedly decreased with cell aging. Control amniotic fluid cell cultures should be derived from cultures for the same serial of time as those from a pregnancy at risk for hereditary lysosomal storage diseases, because the use of later subcultures or other fibroblast cultures as control materials may lead to erroneous interpretations.

CC Genetics - Human 03508

Clinical biochemistry - General methods and applications 10006

Biochemistry studies - Proteins, peptides and amino acids 10064

Biochemistry studies - Carbohydrates 10068

Enzymes - Methods 10804

Enzymes - Physiological studies 10808

Pathology - Diagnostic 12504

Metabolism - Carbohydrates 13004  
 Metabolism - Metabolic disorders 13020  
 Blood - Other body fluids 15010  
 Respiratory system - Physiology and biochemistry 16004  
 Reproductive system - General and methods 16501  
 Bones, joints, fasciae, connective and adipose tissue - Physiology and biochemistry 18004  
 Development and Embryology - Pathology 25503

IT Major Concepts  
 Clinical Chemistry (Allied Medical Sciences); Development; Enzymology  
 (Biochemistry and Molecular Biophysics); Genetics; Metabolism;  
 Pathology; Reproductive System (Reproduction)

IT Miscellaneous Descriptors  
 HUMAN CULTURED AMNIOTIC FLUID CELLS SKIN FIBROBLASTS EMBRYONIC LUNG  
 FIBROBLASTS BETA-D GALACTOSIDASE ALPHA-D MANNOSIDASE BETA  
 HEXOSAMINIDASE BETA-D GLUCURONIDASE ALPHA-D GALACTOSIDASE ALPHA-D  
 GLUCOSIDASE ARYLSULFATASE BETA-D GLUCOSIDASE ALPHA-L FUCOSIDASE ALPHA-D  
 NEURAMINIDASE ALPHA-L IDURONIDASE

ORGN Classifier  
 Hominidae 86215  
 Super Taxa  
 Primates; Mammalia; Vertebrata; Chordata; Animalia  
 Taxa Notes  
 Animals, Chordates, Humans, Mammals, Primates, Vertebrates

RN 9025-42-7 (ALPHA-D-MANNOSIDASE)  
 9012-33-3 (BETA-HEXOSAMINIDASE)  
 9001-45-0 (BETA-D-GLUCURONIDASE)  
 9025-35-8 (ALPHA-D-GALACTOSIDASE)  
 9001-42-7 (ALPHA-D-GLUCOSIDASE)  
 9016-17-5 (ARYLSULFATASE)  
 9001-22-3 (BETA-D-GLUCOSIDASE)  
 9037-65-4 (ALPHA-L-FUCOSIDASE)  
 9001-67-6 (NEURAMINIDASE)  
 9073-56-7 (ALPHA-L-IDURONIDASE)  
 9027-52-5 (BETA HEXOSAMINIDASE)

=> => d all abex tech 124 tot

L24 ANSWER 1 OF 1 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN  
 AN 2005-335264 [35] WPIX  
 DNN N2005-274187 DNC C2005-104144

TI Method of selecting population of women having risk of developing  
 obstetric or gynecologic pathologies e.g. urologic disorders involves  
 determining levels of sialidase and/or prolidase  
 activity and pH value of body fluid sample.

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 IN CAUCI, S  
 PA (UNIS) UNIBIOS SRL  
 CYC 36

PI EP-----1528396 A1 20050504 (200535)\* EN 19 G01N-033-569  
 R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HR HU IE IT LI LT LU  
 LV MC MK NL PL PT RO SE SI SK TR  
 CA-----2485854 A1 20050430 (200535) EN C12Q-001-37 <--  
 US--2005095660 A1 20050505 (200535) C12Q-001-34  
 CN-----1637150 A 20050713 (200576) C12Q-001-25

ADT EP-----1528396 A1 2004EP-0022918 20040927; CA-----2485854 A1  
 2004CA-2485854 20041025; US--2005095660 A1 2003US-0698795 20031031;  
 CN-----1637150 A 2004CN-0080999 20041026

PRAI 2003US-0698795 20031031

IC ICM C12Q-001-25; C12Q-001-34; C12Q-001-37; G01N-033-569  
 ICS G01N-033-48; G01N-033-50; G01N-033-52;  
 G01N-033-84; G06F-017-60

AB EP 1528396 A UPAB: 20050603  
 NOVELTY - Method of selecting population of women having a risk of  
 developing obstetric or gynecologic pathologies involves determining

levels of **sialidase** and/or **prolidase** activity in body fluid sample (f1) by established procedures; determining pH value of (f1); and selecting samples having a **sialidase** value of at least 5.0 nmol of methoxyphenol and/or a **prolidase** level of at least 1500 MOD for **prolidase** and a pH of at least 5.0.

**DETAILED DESCRIPTION** - Method of selecting particular population of women having risk of developing obstetric or gynecologic pathologies as indicated as OR value of at least 5.5 involves determining levels of **sialidase** by procedure described in Cauci et al. Am J Obstet Gynecol 1998; 178; 511-5 and/or **prolidase** activity by procedure described in Cauci et al. J Infect Dis 1998; 178; 1698-706, and pH value of the body fluid samples; and selecting the samples having a **sialidase** value of at least 5.0 nmol of methoxyphenol and/or a **prolidase** level of at least 1500 MOD for **prolidase** and pH of at least 5.0.

**INDEPENDENT CLAIMS** are included for the following:

(1) selecting (M1) a particular population of women having a risk of developing, VLBW, delivery at less than 37 weeks gestation (preferably less than 35 weeks gestation, especially less than 32 weeks gestation) involving: determining levels of **sialidase** as above, and pH value of the body fluid samples; and selecting the samples having a **sialidase** value of at least 0.19 nmol of methoxyphenol and/or a **prolidase** level value of above 22 MOD for **prolidase** and pH of at least 5.0; and

(2) a kit comprising a **sialidase** and/or **prolidase** activity assay in solution that includes a colorless substrate solution in which to inoculate the biologic sample, a developing solution in a container equipped with dispenser, a reference scale to correlate the level of **sialidase** activity of at least 0.19 nmol of methoxyphenol and/or **prolidase** level of at least 22 MOD with the intensity of the developed color, a pH indicator, a reference scale to correlate the pH detected by the indicator with a pH at least 5.0, and an illustrative leaflet containing the instructions for the proper use of the kit.

**USE** - For the determination of the risk of obstetric and gynecologic complications (e.g. low birth weight (LBW), very low birth weight (VLBW), preterm delivery (delivery at less than 37 weeks gestation, PTD), early preterm delivery (delivery at less than 35 or 32 weeks gestation, EPTD), premature rupture of membranes, preterm premature rupture of membranes, intraamniotic infections, spontaneous abortion, endometritis, obstetric surgery infections, post-partum or post-gynecologic surgery infections, pelvic surgery infections, upper genital tract infections which cause infertility, pelvic inflammatory disease (PID), annexitis, cervicitis, sexually transmitted diseases and infections, malignancies of the urogenital tract) in samples of body fluids such as vaginal fluid (claimed).

**ADVANTAGE** - The identification of a threshold of pH greater than or equal to 5.0 in combination with a high **sialidase** and/or **prolidase** activity in body fluid samples is a crucial issue to select woman who have a risk of developing the described pathologies which is found to be 20-30 fold higher than normal woman. The prior art measured pH equal or higher than 4.7. Therefore, a very important selection among women who can develop the pathologies can be put at the attention of the physician. The method is able to predict if the risk is within the 37 weeks gestation or within 35 or even within 32 weeks gestation. It is able to predict the risk of birth of an infant of less than 1500 g, which is associated with severe morbidity and high rate of newborn death; it allows to predict the very high risk even from non-pregnant women just by detecting the **sialidase** and/or **prolidase** activity and pH value; it identifies population of women having a high risk of developing obstetric and gynecologic complications at an early stage of gestation in order to furnish the physician with a valuable tool to decide whether or not to administer a pharmacological therapy. The leaflet correlates the enzymatic activity with the pH value in order to evaluate the risk of pathologies as absent or low (-), medium (+), high (++) or very high (+++).

Dwg.0/0  
 FS CPI EPI  
 FA AB; DCN  
 MC CPI: B04-B04L; B04-L05; B06-A01; B06-D01; B07-D03; B10-A17; B11-C07B1;  
 B11-C07B3; B12-K04A; D05-H09  
 EPI: S03-E09E; S03-E14H2; S03-F10  
 TECH UPTX: 20050603  
 TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Method: (M1) involves selecting the samples having a pH of at least 5, sialidase value of above 0.19 nmol, 0.38 nmol or 2.50 nmol of methoxyphenol and prolidase level of above 22 mOD, 44 mOD, 1000 mOD, 1500 mOD or 2000 mOD. The OR value is calculated and corrected by a standard factor by the SPSS computer statistic program. After the determination of levels of sialidase and/or prolidase activity, phase a score of the levels of sialidase and/or prolidase activity is determined. The pH of the sample is 5 - 7 (preferably 5 - 6, especially 5 - 5.5). The method is carried out in samples of body fluid of pregnant women (preferably women in the first or second trimester of gestation, especially 6 - 24th full week of gestation) or non-pregnant women. Preferred Kit: The pH indicator comprises a revealing paper with a turning interval of 5 - 7 (preferably 5 - 6, especially 5 - 5.5). The reference scale for the sialidase and/or prolidase activity reports standard values associated with enzyme detecting colors. The reference scale for pH value associates the turning interval with a particular color intensity of the same color. The kit includes a test on solid support (preferably on reactive strip or platform test) for the determination of the sialidase and/or prolidase activity. For the determination of sialidase activity, the kit comprises a chromogenic or fluorogenic substrate selected from 2-(3'-methoxyphenyl)-N-acetyl-D-neuraminic acid, 2-O-(o-nitrophenyl)-alpha-D-N-acetyl neuraminic acid, 2'-(4-methylumbelliferyl)-alpha-D-N-acetyl neuraminic acid sodium salt or 5-bromo-4-chloro-3-indolyl-alpha-D-N-acetyl neuraminic acid. For the determination of prolidase activity, the chromogenic or fluorogenic substrate selected is L-proline-para-nitroanilide, L-proline-beta-naphthylamide, N-benzoyloxycarbonyl-L-prolyl-beta-naphthylamide, N-benzoyloxycarbonyl-L-proline-para-nitrophenyl ester, hydroxy-L-prolyl-beta-naphthylamide, L-proline-7-amido-4-methyl-coumarin or L-proline-4-methoxy-beta-naphthylamide.

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(FILE 'HOME' ENTERED AT 07:36:41 ON 03 MAR 2006)

FILE 'REGISTRY' ENTERED AT 07:37:55 ON 03 MAR 2006  
 ACT GIT795F1/A

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L1 (      1)SEA FILE=HCAPLUS ABB=ON  PLU=ON  US2005095660/PN OR US2003-6987
L2      SEL  PLU=ON  L1 1- RN :      13 TERMS
L3 (      13)SEA FILE=REGISTRY ABB=ON  PLU=ON  L2
L4      2 SEA FILE=REGISTRY ABB=ON  PLU=ON  (9001-67-6/BI OR 9025-32-5/BI
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FILE 'HCAPLUS' ENTERED AT 07:38:09 ON 03 MAR 2006  
 ACT GIT795F0/A

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L5 (      549)SEA FILE=HCAPLUS ABB=ON  PLU=ON  DIPEPTIDASE (1A)PROLINE OR PRO
L6 (      13682)SEA FILE=HCAPLUS ABB=ON  PLU=ON  NEURAMINIDASE OR ACETYLNEURAMI
L7      14222 SEA FILE=HCAPLUS ABB=ON  PLU=ON  (L5 OR L6)
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FILE 'BIOSIS' ENTERED AT 07:38:36 ON 03 MAR 2006

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L8      14273 L4,L7
L9      1061 L8 AND (PH OR HYDROGEN (1W)ION)
L10     171 L9 AND ?ASSAY?
L11     165 L10 AND PY<=2003

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E BODY FLUIDS/CT  
L12 0 E3 AND L11  
L13 0 E3 AND L10  
L14 1 L11 AND BODY (1W) FLUID  
L15 6 L11 AND (GYNECOL? OR PREGNAN? OR OBSTET? OR VAGIN?)  
SEL AN 3  
L16 1 L15 AND E1

FILE 'WPIX' ENTERED AT 08:17:48 ON 03 MAR 2006

L17 2187 C12Q001-37/IPC  
L18 682 L7  
E CAUCI S/AU  
L19 3 E3  
L20 54632 G01N033-84/IPC OR (N421 OR N422 OR N425)/M0,M1,M2,M3,M4,M5,M6  
L21 232874 (G01N033-48? OR G01N033-49? OR G01N033-50 OR G01N033-52 OR G01N  
L22 53 L17-18 AND L20  
L23 27 L22 AND L21  
L24 1 L23 AND L19  
L25 26 L23 NOT L24

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